Amendments To The Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

What is claimed is:

(Currently Amended) <u>A method for the treatment of a schizophrenic</u>
 disorder in a mammal in need thereof, said method comprising
 administering to said mammal an effective amount of Use of a compound of formula (I)

$$\mathbb{R}^{2}$$
 \mathbb{N} \mathbb{N} \mathbb{N} \mathbb{N} \mathbb{N} \mathbb{N}

or a pharmaceutically acceptable salt or solvate thereof, in which:

is selected from the group consisting of H, C₁₋₆alkyl, C₁₋₂alkyl substituted by one to five fluorine atoms, C₃₋₆alkenyl, C₃₋₆alkynyl, C₃₋₁₀cycloalkylC₀₋₆alkyl, C₄₋₁₂bridged cycloalkyl, A(CR⁴R⁵)_n and B(CR⁴R⁵)_n;

R² is C₁₋₂alkyl substituted by one to five fluorine atoms;

R³ is selected from the group consisting of C₁₋₆alkyl, NH₂ and

R⁴ and R⁵ are independently selected from H or C₁₋₆alkyl;

R7CONH:

A is selected from the group consisting of unsubstituted 5or 6-membered heteroaryl,unsubstituted 6-membered aryl, 5- or 6-membered heteroaryl substituted by one or more R⁶ and6-membered aryl substituted by one or more R⁶;

R⁶ is selected from the group consisting of halogen, C₁₋
ealkyl, C_{1-ealkyl} substituted by one more fluorine atoms,

C₁₋₆alkoxy, C₁₋₆alkoxy substituted by one or more F, NH₂SO₂ and C₁₋₆alkylSO₂;

B is a ring selected from the group consisting of

where defines the point of attachment of the ring;

R⁷ is selected from the group consisting of H, C_{1-6} alkyl, C_{1-6} alkyl, and C_{1-6} alkyl, C_{1-6} alkyl, and is 0 to 4;

in the preparation of a medicament for the treatment of schizophrenic disorders.

 (Currently Amended) A method for the treatment of a schizophrenic disorder in a mammal in need thereof, said method comprising administering to said mammal an effective amount of Use of a compound of formula (II)

or a pharmaceutically acceptable salt or solvate thereof in which:

 Z^0 is selected from the group consisting of halogen, C_{1-6} alkyl, C_{1-6} alkoxy, C_{1-6} alkoxy substituted by one or more fluorine atoms, and $O(CH_2)_nNZ^4Z^5$;

Z¹ and Z² are each the same or different and are independently selected from the group consisting of H, C₁₋₆alkyl, C₁₋₆alkyl

substituted by one or more fluorine atoms, C_{1-6} alkoxy, C_{1-6} hydroxyalkyl, SC_{1-6} alkyl, C(O)H, $C(O)C_{1-6}$ alkyl, C_{1-6} alkylsulphonyl, C_{1-6} alkoxy substituted by one or more fluorine atoms, $O(CH_2)_nCO_2C_{1-6}$ alkyl, $O(CH_2)_nSC_{1-6}$ alkyl, $O(CH_2)_nNZ^4Z^5$, $O(CH_2)_nSC_{1-6}$ alkyl and $O(C)NZ^4Z^5$; with the proviso that when O(C)0 is at the 4-position and is halogen, then at least one of O(C)1 and O(C)2 is O(C)3 and O(C)4. Salkoxy substituted by one or more fluorine atoms, O(C)5 considerable O(C)6 alkyl, O(C)6 alkyl, O(C)7 alkyl, O(C)8 alkyl, O(C)9 alkyl, O(C

 Z^3 is C_{1-6} alkyl or NH_2 ;

Z⁴ and Z⁵ are each the same or different and are independently selected from the group consisting of H, or C₁₋₆alkyl or, Z⁴ and Z⁵ together with the nitrogen atom to which they are bound, form a 4 - 8 membered saturated heterocyclic ring having 1 or 2 heteroatoms selected from N, O and S; and

n is 1-4; in the preparation of a medicament-for the treatment of schizophrenic disorders.

3. (Currently Amended) A method for the treatment of a schizophrenic disorder in a mammal in need thereof, said method comprising administering to said mammal an effective amount of Use of a compound of formula (III)

$$Q^{10}$$
 Q^{10}
 Q

or a pharmaceutically acceptable salt or solvate thereof in which:

X is selected from the group consisting of oxygen or NQ²;

Y is selected from the group consisting of CH or nitrogen;

is selected from the group consisting of H, C₁₋₆alkyl, C₁₋₂alkyl substituted by one to five fluorine atoms, C₁₋₃alkylOC₁₋₃alkyl, C₃₋₆alkenyl, C₃₋₆alkynyl, C₃₋₁₀cycloalkylC₀₋₆alkyl, C₄₋₇cycloalkyl substituted by C₁₋₃alkyl or C₁₋₃alkoxy, C₄₋₁₂bridged cycloalkyl, A(CR⁶R⁷)_n and B(CR⁶R⁷)_n;

 Q^2 is selected from the group consisting of H and C₁₋₆alkyl; or Q^1 and Q^2 together with the nitrogen atom to which they are bound form a 4-8 membered saturated heterocyclic ring or a 5-membered heteroaryl ring heteroaryl ring is unsubstituted or substituted by one R^8 ;

Q³ is selected from the group consisting of C₁₋₅alkyl and C₁₋₂alkyl substituted by one to five fluorine atoms;

Q⁴ is selected from the group consisting of C₁₋₆alkyl, NH₂ and R⁹CONH;

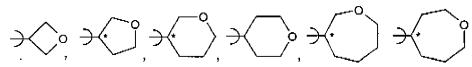
is selected from the group consisting of hydrogen, C_{1-3} alkyl, C_{1-2} alkyl substituted by one to five fluorine atoms, C_{1-3} alkyl O_2 C, halogen, cyano, $(C_{1-3}$ alkyl $)_2$ NCO, C_{1-3} alkyl $)_2$ S;

Q⁶ and Q⁷ are independently H or C₁₋₆alkyl;

A is selected from the group consisting of unsubstituted 5- or 6-membered heteroaryl unsubstituted 6-membered aryl, 5- or 6-membered heteroaryl substituted by one or more R⁸; and 6-membered aryl substituted by one or more R⁸;

is selected from the group consisting of halogen, C₁₋₆alkyl, C₁₋₆alkyl substituted by one more fluorine atoms, C₁₋₆alkoxy, C₁₋₆alkoxy substituted by one or more F, NH₂SO₂ and C₁₋₆alkylSO₂;

B is a ring selected from the group consisting of



and where) defines the point of attachment of the ring;

is selected from the group consisting of H, C₁₋₆alkyl, C₁₋₆alkyl and

C₁₋₆alkylCONHC₁₋₆alkyl;

Q¹⁰ is selected from the group consisting of H and halogen; and n is 0 to 4;

in the preparation of a modicament for the treatment of schizophrenic disorders.

- 4. (Currently Amended) The method of claim 1, further comprising Use of a compound of formula (I), (II) and (III), as defined in anyone of claims from 1 to 3, and pharmaceutically acceptable salts and solvates thereof, administering in combination with a neuroleptic drug in the preparation of a medicament for the treatment of schizophrenic disorders such as schizophrenia, delucional disorders, affective disorders, autism and tic disorders.
- 5. (Canceled)
- 6. (Currently Amended) A method for the treatment of a schizophrenic disorder in a mammal in need thereof, said method comprising administering to said mammal an effective amount of Use of according to Claim 5, wherein the compound is 2-butoxy-4-[4-(methylsulfonyl)phenyl]-6-(trifluoromethyl)pyrimidine or a pharmaceutical acceptable salt or solvate thereof.
- 7. (Currently Amended) The method Use according to Claim 4, characterised in that the neuroleptic is selected from clozapine, olanzapine, ziprasidone, risperidone, aripiprazole, quetiapine, quetiapine fumarate, sertindole, amisulpride, haloperidol, haloperidol decanoate, haloperidol lactate, chlorpromazine, fluphenazine, fluphenazine decanoate, fluphenazine enanthate, fluphenazine hydrochloride,

thiothixene, thiothixene hydrochloride, trifluoperazine, perphenazine, amitriptyline, thioridazine, mesoridazine, molindone, molindone hydrochloride, loxapine, loxapine hydrochloride, loxapine succinate, pimozide, flupenthixol, promazine, triflupromazine, chlorprothixene, droperidol, actophenazine, prochlorperazine, methotrimeprazine, pipotiazine, ziprasidone, hoperidone, zuclopenthixol, and mixtures thereof.

- 8. (Currently Amended) The method Use according to Claim 4, wherein the neuroleptic is risperidone or aripiprazole.
- 9. (Currently Amended) The method of claim 1 wherein the compound is Use of 2-butoxy-4-[4-(methylsulfonyl)phenyl]-6-(trifluoromethyl)pyrimidine or a pharmaceutical acceptable salt thereof, <u>further comprising</u> in combination with risperidone in an amount of 0.8-3.0 mg/kg and 2-6 mg, respectively, in the preparation of a medicament for the treatment of schizophrenic disorders such as schizophrenia, delusional disorders, affective disorders, autism and tic disorders.
- (Currently Amended) <u>The method of claim Use according to Claim 9</u>,
 wherein risperidone is administered in an amount of 4-5 mg.
- 11. (Canceled)
- 12. (Currently Amended) Kit-of-parts suitable for use in the treatment of schizophrenic disorders such as schizophrenia, delusional disorders, affective disorders, autism or tic disorders, schizophreniform disorders, in particular chronic schizophrenic psychoses and schizoaffective psychoses, temporary acute psychotic disorders, comprising a first dosage form comprising a neuroleptic drug and a second dosage form comprising a compound of formula (I) (II) and (III) as defined in anyone of claim from 1 to 3 or a pharmaceutical acceptable salt or solvate thereof, for simultaneous, separate or sequential administration.

- 13. (Original) Kit-of-parts according to Claim 12, characterised in that the neuroleptic is selected from the group consisting of: clozapine, olanzapine, ziprasidone, risperidone, quetiapine, quetiapine fumarate, sertindole, amisulpride, haloperidol, haloperidol decanoate, haloperidol lactate, chlorpromazine, fluphenazine, fluphenazine decanoate, fluphenazine enanthate, fluphenazine hydrochloride, thiothixene, thiothixene hydrochloride, trifluoperazine, perphenazine, amitriptyline, thioridazine, mesoridazine, molindone, molindone hydrochloride, loxapine, loxapine hydrochloride, loxapine succinate, pimozide, flupenthixol, promazine, triflupromazine, chlorprothixene, droperidol, actophenazine, prochlorperazine, methotrimeprazine, pipotiazine, ziprasidone, hoperidone, zuclopenthixol, and mixtures thereof.
- 14. (Currently Amended) Kit-of-parts according to claims-12 and 13, further comprising a compound selected from the group consisting of: celecoxib, rofecoxib, meloxicam, piroxicam, deracoxib, parecoxib, valdecoxib, etoricoxib, a chromene derivative, a chroman derivative, N- (2-cyclohexyloxynitrophenyl) methyl sulfonamide, COX189, ABT963 or JTE-522, or pharmaceutical acceptable salts or solvates thereof.
- 15. (Currently Amended) Kit-of-parts according to <u>claim 12</u> anyone of <u>Claims</u> from 12 to 14, wherein said compound is 2-butoxy-4-[4- (methylsulfonyl)phenyl]-6-(trifluoromethyl)pyrimidine or a pharmaceutical acceptable salt thereof and said neuroleptic drug is risperidone.
- 16. (Original) Kit-of-parts according to Claim 15, wherein 2-butoxy-4-[4-(methylsulfonyl)phenyl]-6-(trifluoromethyl)pyrimidine or a pharmaceutical acceptable salt thereof and risperidone are in an amount of 0.8-3.0 mg/kg mg and 2-6 mg, respectively.
- 17. (Canceled)

- 18. (Currently Amended) The method according to claim <u>1</u> 17, wherein said mammal is human.
- 19. (Original) The method according to claim 18, wherein said schizophrenic disorder is selected from the group consisting of schizophrenia, delusional disorders, affective disorders, autism or tic disorders, schizophreniform disorders, in particular chronic schizophrenic psychoses and schizoaffective psychoses, temporary acute psychotic disorders.
- 20. (Original) The method according to claim 19, further comprising administering a therapeutically effective amount of a a neuroleptic drug.
- 21. (Original) The method according to claim 20, wherein said neuroleptic drug is selected from the group consisting of: clozapine, olanzapine, ziprasidone, risperidone, quetiapine, quetiapine fumarate, sertindole, amisulpride, haloperidol, haloperidol decanoate, haloperidol lactate, chlorpromazine, fluphenazine, fluphenazine decanoate, fluphenazine enanthate, fluphenazine hydrochloride, thiothixene, thiothixene hydrochloride, trifluoperazine, perphenazine, amitriptyline, thioridazine, mesoridazine, molindone, molindone hydrochloride, loxapine, loxapine hydrochloride, loxapine succinate, pimozide, flupenthixol, promazine, triflupromazine, chlorprothixene, droperidol, actophenazine, prochlorperazine, methotrimeprazine, pipotiazine, ziprasidone, hoperidone, zuclopenthixol, and mixtures thereof.
- 22. (Original) A method for the treatment of a schizophrenic disorder in a mammal in need thereof, said method comprising administering to said mammal a therapeutically effective amount of a compound, the compound is selected from the group consisting of: 2-(4-fluorophenoxy)-4-[4-(methylsulfonyl)phenyl]-6](trifluoromethyl)pyrimidine;

- 2-(4-methoxyphenoxy)-4-[4-(methylsulfonyl)phenyl]-6trifluoromethyl)pyrimidine;
- 2-butoxy-4-[4-(methylsulfonyl)phenyl]-6-(trifluoromethyl)pyrimidine;
- 2-[(5-chloropyridin-3-yl)oxy]-4-[4-(methylsulfony)phenyl]-6-(trifluoromethyl)pyrimidine;
- 2-(cyclohexyloxy)-4-[4-(methylsulfonyl)phenyl]-6-(trifluoromethyl)pyrimidine;
- 3-(4-methylsulfonyl-phenyl)-2-(4-methoxy-phenyl)-pyrazolo[1,5-b]pyridazine;
- 6-difluoromethoxy-2-(4-fluoro-phenyl)-3-(4-methylsulfonyl-phenyl)-pyrazolo[1,5-b]-pyridazine;
- 2-(4-ethoxy-phenyl)-3-(4-methylsulfonyl-phenyl)-pyrazolo[1,5-b]pyridazine;
- 2-(4-fluoro-phenyl)-6-methylsulfonyl-3-(4-methylsulfonyl-phenyl)-pyrazolo[1,5-b]pyridazine;
- 2-(4-difluoromethoxy-phenyl)-3-(4-methylsulfonyl-phenyl)-pyrazolo[1,5-b]pyridazine;
- 4-[2-(4-ethoxy-phenyl)-pyrazolo[1,5-b]pyridazin-3-yl]-benzenesulfonamide;
- 6-difluoromethoxy-2-(3-fluoro-phenyl)-3-(4-methylsulfonyl-phenyl)-pyrazolo[1,5-b]pyridazine;
- 3-(4-methanesulfonyl-phenyl)-2-(4-methoxy-phenyl)-pyrazolo[1,5-b]pyridazine;
- 6-difluoromethoxy-2-(4-fluoro-phenyl)-3-(4-methanesulfonyl-phenyl)-pyrazolo[1,5-b]pyridazine;
- 2-(4-ethoxy-phenyl)-3-(4-methanesulfonyl-phenyl)-pyrazolo[1,5-b]pyridazine;
- 2-(4-fluoro-phenyl)-6-methanesulfonyl-3-(4-methanesulfonyl-phenyl)-pyrazolo[1,5-b]pyridazine;
- 2-(4-difluoromethoxy-phenyl)-3-(4-methanesulfonyl-phenyl)-pyrazolo[1,5-b]pyridazine;
- 4-[2-(4-ethoxy-phenyl)-pyrazolo[1,5-b]pyridazin-3-yl]-benzenesulfonamide;

6-difluoromethoxy-2-(3-fluoro-phenyl)-3-(4-methanesulfonyl-phenyl)-pyrazolo[1,5-b]pyridazine

4-ethyl-6-[4-(methylsulfonyl)phenyl]-N-(tetrahydro-2H-pyran-4-ylmethyl)-2-pyridinamine;4-methyl-N-[(1-methyl-1H-pyrazol-4-yl)methyl]-6-[4-(methylsulfonyl)phenyl]-2-pyridinamine;

N-[(1,5-dimethyl-1H-pyrazol-4-yl)methyl]-4-methyl-6-[4-(methylsulfonyl)phenyl]-2-pyridinamine;

N-[(1,3-dimethyl-1H-pyrazol-4-yl)methyl]-4-methyl-6-[4-(methylsulfonyl)phenyl]-2-pyridinamine;

4-(6-{[(1,3-dimethyl-1H-pyrazol-4-yl)methyl]amino}-4-ethyl-2-pyridinyl)benzenesulfonamide;

N-[(1,3-dimethyl-1H-pyrazol-4-yl)methyl]-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

N-[(1,5-dimethyl-1H-pyrazol-4-yl)methyl]-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

4-{4-methyl-6-[(tetrahydro-2H-pyran-4-ylmethyl)amino]-2-pyridinyl}benzenesulfonamide;

4-methyl-N-[(1-methyl-1H-pyrazol-3-yl)methyl]-6-[4-(methylsulfonyl)phenyl]-2-pyridinamine;

N-(cyclohexylmethyl)-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

N-cyclohexyl-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

2-[4-(methylsulfonyl)phenyl]-6-[(2-pyridinylmethyl)oxy]-4-(trifluoromethyl)pyridine;

4-methyl-N-[(3-methyl-4-isoxazolyl)methyl]-6-[4-(methylsulfonyl)phenyl]-2-pyridinamine;

6-[4-(methylsulfonyl)phenyl]-N-(2-pyridinylmethyl)-4-(trifluoromethyl)-2-pyridinamine;

N-cycloheptyl-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

N-(cis-4-methylcyclohexyl)-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

N-(1-ethylpropyl)-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

N-[(3-methyl-1,2,4-oxadiazol-5-yl)methyl]-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

N-[(5-methyl-1,2,4-oxadiazol-3-yl)methyl]-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

4-methyl-N-[(1-methyl-1H-pyrazol-5-yl)methyl]-6-[4-(methylsulfonyl)phenyl]-2-pyridinamine;

N-(cyclopentylmethyl)-6-[4-(methylsulfonyl)phenyl]-4-(trifluoromethyl)-2-pyridinamine;

N-[(1-ethyl-1H-1,2,4-triazol-5-yl)methyl]-4-methyl-6-[4-(methylsulfonyl)phenyl]-2-pyridinamine;

4-ethyl-6-[4-(methylsulfonyl)phenyl]-2-[(2-pyridinylmethyl)amino]-3-pyridinecarbonitrile;

4-ethyl-2-{[(5-methyl-2-pyridinyl)methyl]amino}-6-[4-

(methylsulfonyl)phenyl]-3-pyridinecarbonitrile;

4-ethyl-2-{[(6-methyl-3-pyridinyl)methyl]amino}-6-[4-

(methylsulfonyl)phenyl]-3-pyridinecarbonitrile;

4-ethyl-2-{[(1-methyl-1H-pyrazol-4-yl)methyl]amino}-6-[4-

(methylsulfonyl)phenyl]-3-pyridinecarbonitrile;

4-ethyl-6-[4-(methylsulfonyl)phenyl]-2-{[(4-methyl-1,3-thiazol-2-yl)methyl]amino}-3-pyridinecarbonitrile;

4-ethyl-6-[4-(methylsulfonyl)phenyl]-2-[(2-pyridinylmethyl)oxy]-3-pyridinecarbonitrile;

4-ethyl-N-[(1-ethyl-1H-1,2,4-triazol-5-yl)methyl]-6-[4-(methylsulfonyl)phenyl]-2-pyridinamine;

4-ethyl-2-{[(6-methyl-3-pyridinyl)methyl]oxy}-6-[4-(methylsulfonyl)phenyl]-3-pyridinecarbonitrile;

6-[4-(methylsulfonyl)phenyl]-N-[(1-methyl-1H-1,2,4-triazol-5-yl)methyl]-4-(trifluoromethyl)-2-pyridinamine; and pharmaceutically acceptable salts and solvates thereof

- 23. (Original) The method according to claim 22, wherein said mammal is human.
- 24. (Original) The method according to claim 23, wherein said schizophrenic disorder is selected from the group consisting of schizophrenia, delusional disorders, affective disorders, autism or tic disorders, schizophreniform disorders, in particular chronic schizophrenic psychoses and schizoaffective psychoses, temporary acute psychotic disorders.
- 25. (Original) The method according to claim 24, further comprising administering a therapeutically effective amount of a a neuroleptic drug.
- 26. (Original) The method according to claim 25, wherein said neuroleptic drug is selected from the group consisting of: clozapine, olanzapine, ziprasidone, risperidone, quetiapine, quetiapine fumarate, sertindole, amisulpride, haloperidol, haloperidol decanoate, haloperidol lactate, chlorpromazine, fluphenazine, fluphenazine decanoate, fluphenazine enanthate, fluphenazine hydrochloride, thiothixene, thiothixene hydrochloride, trifluoperazine, perphenazine, amitriptyline, thioridazine, mesoridazine, molindone, molindone hydrochloride, loxapine, loxapine hydrochloride, loxapine succinate, pimozide, flupenthixol, promazine, triflupromazine, chlorprothixene, droperidol, actophenazine, prochlorperazine, methotrimeprazine, pipotiazine, ziprasidone, hoperidone, zuclopenthixol, and mixtures thereof.
- 27. (Original) A method for the treatment of a schizophrenic disorder in a mammal in need thereof, said method comprising administering to said mammal a therapeutically effective amount of 2-butoxy-4-[4-(methylsulfonyl)phenyl]-6-(trifluoromethyl)pyrimidine or a pharmaceutical acceptable salt or solvate thereof.

- 28. (Original) The method according to claim 27, wherein said mammal is human.
- 29. (Original) The method according to claim 28, wherein said schizophrenic disorder is selected from the group consisting of schizophrenia, delusional disorders, affective disorders, autism or tic disorders, schizophreniform disorders, in particular chronic schizophrenic psychoses and schizoaffective psychoses, temporary acute psychotic disorders.
- 30. (Original) The method according to claim 28, further comprising administering a therapeutically effective amount of a a neuroleptic drug.
- 31. (Original) The method according to claim 30, wherein said neuroleptic drug is selected from the group consisting of: clozapine, olanzapine, ziprasidone, risperidone, quetiapine, quetiapine fumarate, sertindole, amisulpride, haloperidol, haloperidol decanoate, haloperidol lactate, chlorpromazine, fluphenazine, fluphenazine decanoate, fluphenazine enanthate, fluphenazine hydrochloride, thiothixene, thiothixene hydrochloride, trifluoperazine, perphenazine, amitriptyline, thioridazine, mesoridazine, molindone, molindone hydrochloride, loxapine, loxapine hydrochloride, loxapine succinate, pimozide, flupenthixol, promazine, triflupromazine, chlorprothixene, droperidol, actophenazine, prochlorperazine, methotrimeprazine, pipotiazine, ziprasidone, hoperidone, zuclopenthixol, and mixtures thereof.
- 32. (Original) The method according to claim 31, wherein said compound is 2-butoxy-4-[4-(methylsulfonyl)phenyl]-6- (trifluoromethyl)pyrimidine or a pharmaceutical acceptable salt thereof and said neuroleptic drug is risperidone.

33. (Original) The method according to claim 32, wherein said compound and said neuroleptic drug are present in an amount of 0.8-3.0 mg/kg mg and 2-6 mg respectively.